

STANFORD

School of Humanities and Sciences



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Science

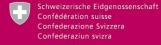
Director, Gendered Innovations in Science, Health & Medicine, Engineering, and Environment















Gendered in Science, Health & Medicine, Innovations Engineering, and Environment

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ENVIRONMENT

What is Gendered Innovations?

SEX & GENDER ANALYSIS

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Paradigm



Why Gendered Innovations?

MEDICINE

"Gendered Innovations" employs methods of sex and gender analysis to create new knowledge.

Percentage of EC Horizon 2020 grants funded in 2015 that integrated sex or gender analysis into research:

36%

European Commission. Horizon 2020. Monitoring Report 2015. Directorate-General for Research and Innovation (2016), 53. (6,062 grants analyzed).

Gendered Innovations...

Can we harness the creative power of sex & gender analysis for discovery?

THREE FIXES

- 1. FIX THE NUMBERS OF WOMEN
- 2. FIX THE INSTITUTIONS
- 3. FIX THE KNOWLEDGE

FIX THE KNOWLEDGE

Doing Research Wrong Costs Lives and Money

Between 1997 and 2000, 10 drugs were withdrawn from the U.S. market because of life-threatening health effects—8 of those showed greater severity in women (GAO, 2001).

Doing Research Right Saves Lives and Money

WHI Hormone Therapy Trials

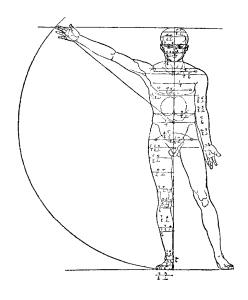
- Each \$1 spent returned \$140 to US taxpayers in health care savings
- Health Improvements
 - * 76,000 fewer cases of cardiovascular disease
 - * 126,000 fewer breast cancer cases
 - * 145,000 more quality-adjusted life years
 - * However: 263,000 more osteoporotic fractures

Joshua A. Roth et al., Annals of Internal Medicine (2014) 160, 9:594-602.

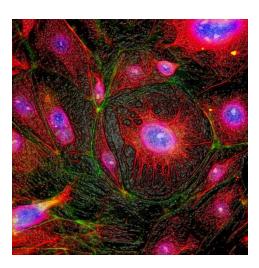
Gendered Innovations

- 1) develop state-of-the-art Methods of sex and gender analysis
- 2) provide Case Studies to illustrate how gender analysis leads to discovery and innovation.

Most research is done in males

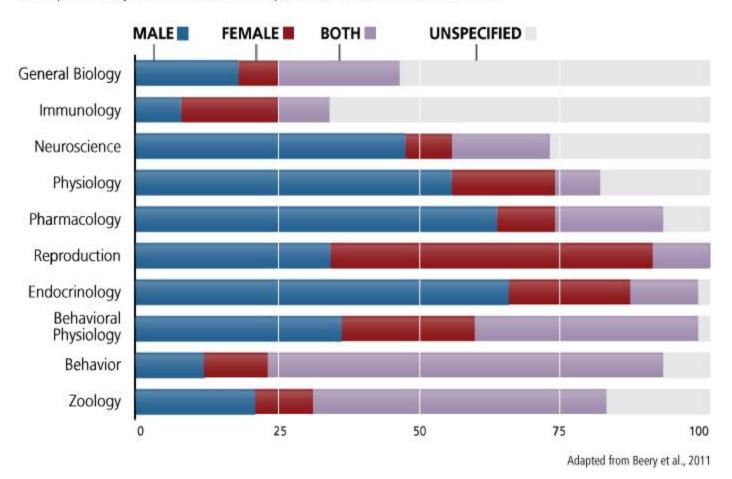






Proportion of Research Studies Using Male and/or Female Animals

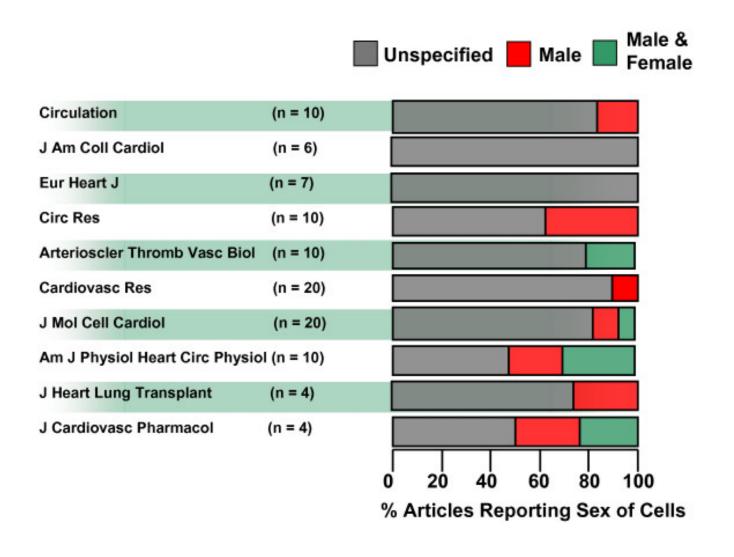
From published journal articles within specified biomedical subfield, 2009



Beery, A., & Zucker, I. (2011). Sex Bias in Neuroscience and Biomedical Research. *Neuroscience and Biobehavioral Reviews, 35 (3)*, 565-572.

Taylor, K., Vallejo-Giraldo, C., Schaible, N., Zakeri, R., & Miller, V. (2011). Reporting of Sex as a Variable in Cardiovascular Studies using Cultured Cells. *Biology of Sex Differences*, *2* (11), 1-7.

Percentage of articles reporting sex of cells used in the experiments



Shah, K., McCormack, C. E., & Bradbury, N. A. (2014). **Do you know the sex of your cells?** *American Journal of Physiology-Cell Physiology*, *306*(1), C3-C18.

Cell Line	Sex	Description	Species	Year	Origin	Reference
5637	Male	Urinary bladder epithelium	Human	1974	С	(68, 69)
3T3-L1	Male	Embryo fibroblast	Mouse	1962	N	(84, 85)
16HBE		Lung epithelial	Human		N	(46, 88)
A549	Male	Lung epithelial	Human	1972	С	(79, 137)
A6	Male	Kidney epithelial	Xenopus	1965	N	(165,222)
A7r5		Aorta smooth muscle	Rat	1976	N	(15, 183)
AGS	Female	Stomach epithelial	Human	1979	С	(10, 11)
AML-12	Male	Liver epithelial	Mouse	1994	N	(61, 282)
AML-193	Female	Lymphoblast	Human	1987	С	(131,229)
ARPE-19	Male	Retinal pigmented epithelial	Human	1986	N	(62, 101)
BeWo	Female	Placenta	Human	1966	N	(193,194)
внк		Kidney fibroblast	Syrian hamster	1961	N	(146,148)

15-Year Outcomes for Heart Transplant

Years post- HT	MD-MR (n=40,497) Mean Survival %	FD-MR (n=13,480) Mean Survival %	MD-FR (n=6,436) Mean Survival %	FD-FR (n=7,420) Mean Survival %
1	83.74	78.95	82.94	81.92
5	67.18	62.49	68.94	68.83
10	51.97	47.53	51.79	52.08
15	33.19	29.63	35.37	<u>37.05</u>

Best outcomes: female-to-female and male-to-male

Kaczmarek et al., "Gender Does Matter: Gender-Specific Outcome Analysis of 67,855 Heart Transplants," *Thorac Cardiovasc Surg*, 2013.

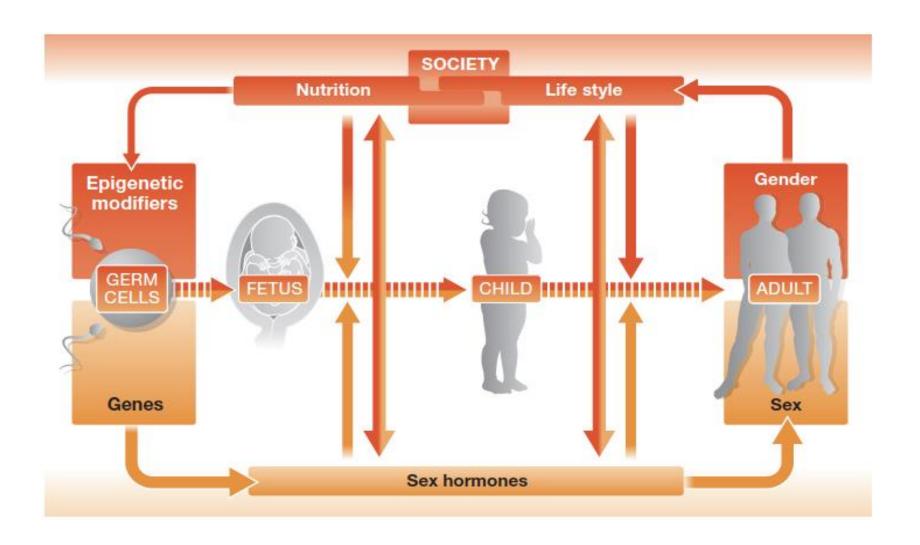
Definition of Terms

SEX = biological characteristics

GENDER = cultural attitudes and behaviors

Sex and Gender Interact

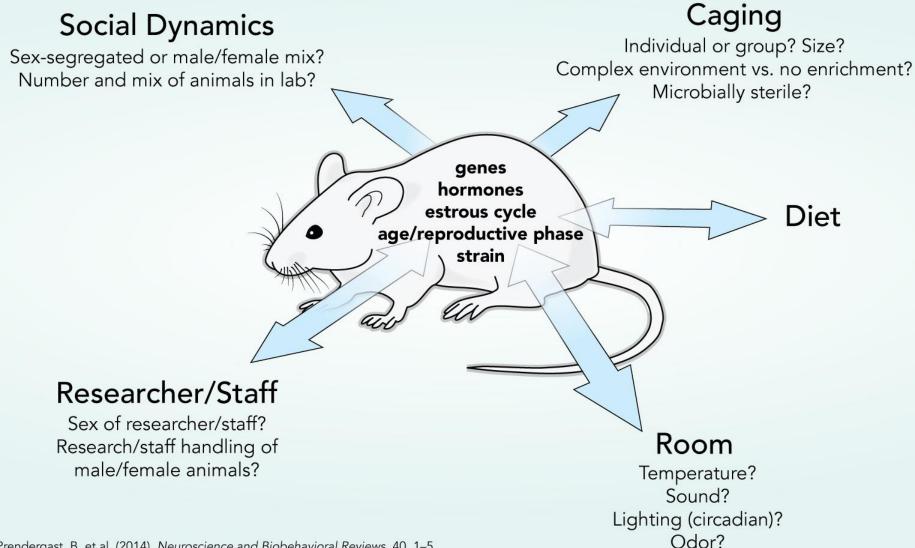
Regitz-Zagrosek, V. (2012). Sex and Gender Differences in Health. EMBO Reports, 13 (7): 596-603.



DANGER!

- That PIs confound sex with gender
- That PIs "see" sex (or a biological trait), when they are really looking at "gender" (or an environmental condition that may impact male and female animals differently).
- Schiebinger, Londa, Seth S. Leopold, and Virginia M. Miller. "Editorial policies for sex and gender analysis." *The Lancet* 388.10062 (2016): 2841-2842.

Integrating Sex & Gender into Animal Research



Prendergast, B. et al. (2014). Neuroscience and Biobehavioral Reviews. 40, 1–5. Ritz, S. et al., (2014). FASEB J, 28(1), 4-13. Sorge, R. et al. (2014). Nature Methods.

Cost?

From the NSF-funded Stanford workshop on Animal Research: Klein, S. L., Schiebinger, L., Stefanick, M. L., Cahill, L., Danska, J., de Vries, G. J., ... & Zucker, I. (2015). Opinion: Sex inclusion in basic research drives discovery. *Proceedings of the National Academy of Sciences*, 112(17), 5257-5258.

Gender in Heart Disease

Can we reduce gender to variables so that results can be manipulated statistically?

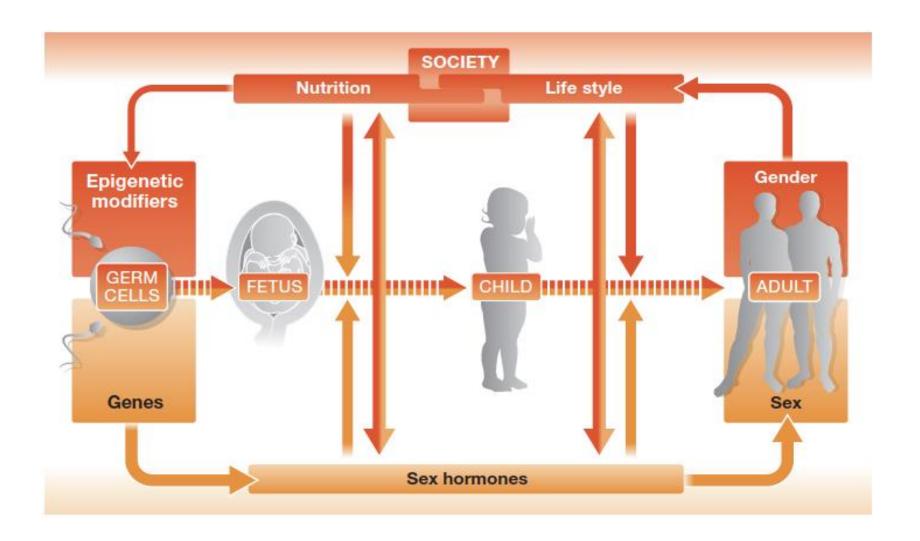
Gender Variables in Health Research

Inspiration from Canada

Pelletier, R., Khan, N. A., Cox, J., Daskalopoulou, S. S., Eisenberg, M. J., Bacon, S. L., ... & Norris, C. M. (2016). Sex versus Gender-Related Characteristics: Which Predicts Outcome After Acute Coronary Syndrome in the Young?. *Journal of the American College of Cardiology*, 67 (2), 127-135.

Sex and Gender Interact

Regitz-Zagrosek, V. (2012). Sex and Gender Differences in Health. EMBO Reports, 13 (7): 596-603.



Seven Relevant Gender Factors

- household primary earner
- > personal income
- number of hours per week spent doing housework
- status of primary person responsible for doing housework
- > level of stress at home
- Bem Sex Role Inventory masculinity score
- Bem Sex Role Inventory femininity score

Acute Coronary Syndrome (aged 55 and younger)

- No sex difference
- Gender matters. Patients received a "genderrelated score" of between 1 and 100. Patients with a higher "femininity" score—regardless whether they were men or women—were more likely to experience a recurrence of ACS.

No longer valid

- household primary earner
- > personal income
- number of hours per week spent doing housework
- status of primary person responsible for doing housework
- > level of stress at home
- Bem Sex Role Inventory masculinity score
- Bem Sex Role Inventory femininity score

Stanford Study

Gender Variables in Health Research

Hoping for results in July

Londa Schiebinger, Mathias Wullum Nielsen, Claudette Brooks, Mark R. Cullen, Gillian Einstein, John P.A. Ioannidis, Ineke Klinge, Hannah LeBlanc, Torsten B. Neilands, Hee Young Paik, Diana Peragine, Louise Pilote, Judith Prochaska, Stephen Ristvedt, Kierstyn Smith, YoonJu Song, Marcia L. Stefanick

Seed Grants:

One example: http://wsdm.stanford.edu/

Research Resources:

- Gendered Innovations website. genderedinnovations.stanford.edu
- March 2015: The EC Gender Advisory Group published an advice paper on preparing grants that integrate the gender dimension into research. http://ec.europa.eu/transparency/regexpert/index.cfm?do=groupDetail.groupDetailDoc&id=18892&no=1
- August 2015: The Canadian Institutes for Health Research rolled out the first of three trainings for Sex and Gender in Biomedical Research.
 http://www.discoversexandgender.ca/

Research & Teaching Resources:

- Oertelt-Prigione, S.& Regitz-Zagrosek, V. (Eds.)
 (2012). Sex and Gender Aspects in Clinical Medicine.
- Regitz-Zagrosek, V., (Ed.) (2012). Sex and Gender Differences in Pharmacology.
- Schenck-Gustafsson, K., DeCola, P., Pfaff, D., Pisetsky, P. (Eds.) (2012). Handbook of Clinical Gender Medicine.

Curriculum

- Sex and Gender-Based Medical Education Summit -A Roadmap for Curricular Innovation, Mayo Clinic, US, October 2015 https://ce.mayo.edu/special-topics-in-health-care/node/3095
- Charité, Berlin: Ludwig, S., et al. (2015). A successful strategy to integrate sex and gender medicine into a newly developed medical curriculum. *Journal of Women's Health*, 24(12), 996-1005.

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On the Gendered Innovations website, sign up at "Contact Us."



Innovative Approaches

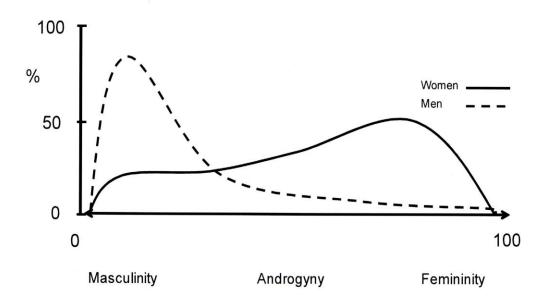
1. To move beyond binary notions of "masculinity" and "femininity."

2. To capture three dimensions of gender:

Norms Identity

Relations

APPENDIX C. Gender-related score distribution in men and women with premature acute coronary syndrome



Proposed Gender-Related Variables

Norms

- Responsibility for Caregiving
- Time use
- Occupation

Identity

- Competitive
- Independent
- Risk-taking
- Communal
- Expressive

Relations

- Social support
- Perceived discrimination
- Quality of family relationships

Communal

- How often do you worry about what other people think about you?
- When making an important decision in your personal life, how often do you take other peoples' needs into account?
- How often do friends talk to you about their problems?
- How easy is it for you to spot when someone in a group is feeling uncomfortable?

Physician Gender Attitudes Interact with Patient Sex and Gender

